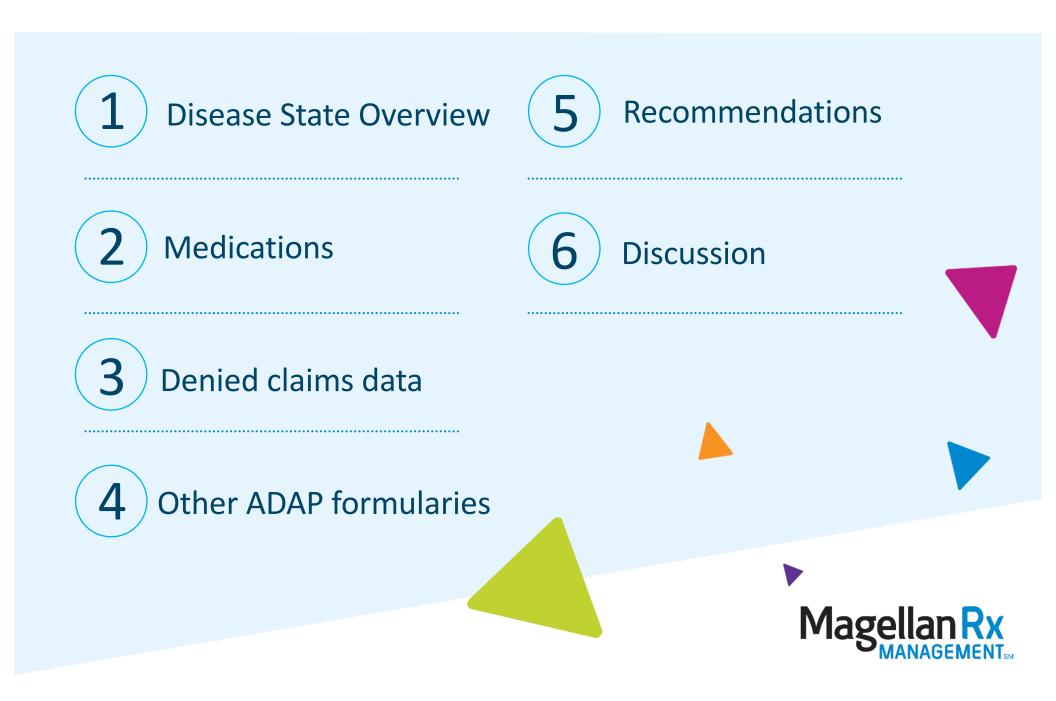


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### **CA-ADAP Formulary Review: AGENDA**



## Therapeutic Class Review

LONG-ACTING INJECTABLE ANTIPSYCHOTIC AGENTS



### Features of Schizophrenia



- **Schizophrenia:** A chronic brain disorder where people interpret reality abnormally. It affects how a person thinks, feels, and behaves.
  - Positive symptoms:
    - Hallucinations
    - Delusions
    - Paranoia

 Exaggerated or distorted perceptions, beliefs, and behaviors

- Negative symptoms:
  - Flattened affect
  - Reduced ability to experience pleasure
- Decreased desire to initiate plans

- Disorganized symptoms:
  - Confused and disordered
    Bizarre behavior thinking and speech
    - Abnormal movements
  - Trouble with logical thinking
- Cognition can also be affected, leading to problems with attention, concentration and memory, and to declining educational performance



## Schizophrenia and Co-Morbid Conditions



- Other conditions, including HIV infection, are more frequent in people with serious mental illness
- Mortality is increased in individuals with schizophrenia, and the average life span is shortened by a decade or more (typically related to comorbid conditions)
- Mental health is a strong predictor of adherence to meds
  - Managing mental health can improve quality of life and psychological wellbeing, therefore increasing adherence to ARV therapy



### Treatment of Schizophrenia



- Schizophrenia requires lifelong treatment, even when symptoms have subsided
  - Goals of Treatment: symptom remission and restoring baseline function at the lowest possible dose
- General guidelines, particularly for patients with symptomatic HIV disease, include the following:
  - Using lower starting doses and slower titration
  - Providing the least complicated dosing schedules possible
  - Focusing on drug side effect profiles to avoid unnecessary adverse events
  - Maintaining awareness of drug metabolism/clearance pathways to minimize drugdrug interactions and possible end organ damage.
- Clinical guidelines suggest treatment with a single agent but do not recommend any agent over another
  - Treatment with more than one antipsychotic should be avoided



## Long-Acting Injectables (LAIs)



- The American Psychiatric Association (APA) guidelines suggests LAI if:
  - Patient preference
  - Non-Adherence history

#### LAIs may be preferred for:

- Difficulty swallowing pills
- Ambivalence
- Inconsistent in swallowing
- Difficulty remembering

#### LAIs may be warranted when:

- Not responding to oral agent
- Transitioning between settings
- Early in schizophrenia treatment
- Known substance abuse disorder



## Long-Acting Injectables (LAIs) cont'd



#### Benefits:

- Decreased risk of mortality
- Reduced risk of hospitalization
- Decreased rates of discontinuation
- ADRs may be less problematic
- Sense of better control
- Better adherence
- Greater convenience

#### Harms:

- ADRs may last longer Injection-related ADRs
- Longer time to steady-state



## Long-Acting Injectable − 1<sup>st</sup> Generation ▶



- Mechanism of Action: Dopamine receptor antagonists
- <u>Place in Therapy</u>: **Not** recommended as initial therapy for most patients
- <u>Side effects</u>: High incidence of EPS, sedation, cardiac abnormalities, hyperprolactinemia, neuroleptic malignant syndrome (NMS)

AGENT	GENERIC	СТ	тх	WA	STRENGTHS
Fluphenazine (PROLIXIN)	Y	-	-	-	•2.5mg, 5mg, 25mg
Haloperidol (HALDOL)	Y	-	-	-	•50mg, 100mg



## Long-Acting Injectable – 2<sup>nd</sup> Generation▶



- Mechanism of Action: Serotonin-dopamine antagonists
- <u>Place in Therapy:</u> Recommended as *initial therapy* for most patients
- <u>Side effects</u>: weight gain and related metabolic effects, hypotension, sedation, anticholinergic symptoms, hyperprolactinemia, EPS, cardiac effects, and sexual dysfunction

AGENT	GENERIC	СТ	ТХ	WA	STRENGTHS
Aripiprazole (ABILIFY MAINTENA)	-	-	-	-	•300mg, 400mg
Aripiprazole lauroxil (ARISTADA)	-	-	-	-	•441mg, 662mg, 882mg, 1064mg, 675mg (Initio)
Olanzapine pamoate (ZYPREXA RELPREVV)	-	-	-	-	•210mg, 300mg, 405mg
Paliperidone (INVEGA SUSTENNA, INVEGA TRINZA, INVEGA HAFYERA)	-	-	-	-	•39mg, 78mg, 117mg, 156mg, 234mg, 273mg, 410mg, 546mg, 819mg
Risperidone (RISPERDAL CONSTA)	-	-	-	-	•12.5mg, 25mg, 37.5mg, 50mg



## Long-Acting Injectable Rejected Claims



AGENT	REJECTIONS
Olanzapine pamoate (ZYPREXA RELPREVV)	-
Risperidone (RISPERDAL CONSTA)	1.5%
Fluphenazine (PROLIXIN)	3%
Haloperidol (HALDOL)	8.5%
Aripiprazole (ABILIFY MAINTENA)	18%
Aripiprazole lauroxil (ARISTADA)	20%
Paliperidone (INVEGA SUSTENNA, INVEGA TRINZA)	49%





# Formulary Management Recommendations: Schizophrenia

Paliperidone (INVEGA)

Recommend addition to the ADAP formulary

Aripiprazole (ABILIFY MAINTENA)

Consider addition to the ADAP formulary

Aripiprazole Lauroxil (ARISTADA)

Consider addition to the ADAP formulary



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